

# **Anticoagulants Classification**

# 1. Parenteral Anticoagulants

#### A. Indirect Anticoagulants

#### 1. Unfractionated Heparin (UFH)

- Indications:
  - Treatment and prevention of venous thromboembolism (VTE)
  - Acute coronary syndrome (ACS)
  - o Dialysis
- Contraindications:
  - Active bleeding
  - Severe thrombocytopenia (low platelet count)
  - History of heparin-induced thrombocytopenia (HIT)
- Notes:
  - Requires monitoring with activated partial thromboplastin time (aPTT).

- Rapid onset and reversal possible with protamine sulfate.
- 2. Low Molecular Weight Heparins (LMWH)
  - **Examples**: Enoxaparin (Lovenox), Dalteparin (Fragmin)
  - Indications:
    - Treatment of deep vein thrombosis (DVT) and pulmonary embolism (PE)
    - Prophylaxis in patients undergoing surgery or immobility
    - Managing ACS
  - Contraindications:
    - Active bleeding
    - Severe renal impairment (caution with dosages)
    - History of HIT
  - Notes:
    - More predictable anticoagulant response than UFH.
    - Usually does not require routine monitoring.
- 3. Fondaparinux
  - Indications:
    - Prophylaxis of DVT in post-surgical patients
    - Treatment of DVT and PE (in combination with warfarin)
    - Acute coronary syndrome

### Contraindications:

- Active bleeding
- Severe renal impairment
- Body weight < 50 kg (caution due to risk of insufficient dosing)
- Notes:
  - Does not require monitoring.
  - Reversal is challenging; usually only supportive measures.

#### **B. Direct Anticoagulants**

- 1. Thrombin Inhibitors
  - **Examples**: Bivalirudin, Dabitran, Argatroban
  - Indications:
    - Bivalirudin: Used in percutaneous coronary interventions (PCI).

• Argatroban: Used in patients with HIT or those undergoing PCI.

# Contraindications:

- Active bleeding
- Severe renal impairment (particularly with dabigatran)
- Notes:
  - Bivalirudin has a low risk of bleeding compared to UFH.
- 2. Factor Xa Inhibitors
  - **Example**: Otamixaban (investigational), Edoxaban, Rivaroxaban, Apixaban
  - Indications:
    - Rivaroxaban and Apixaban: Prevention and treatment of DVT and PE, stroke prevention in atrial fibrillation.
    - Edoxaban: Similar indications as above.
    - Otamixaban: Under study for ACS.
  - Contraindications:
    - Active bleeding
    - Severe renal disease (caution advised with all except when dosing is adjusted)
  - Notes:
    - Minimal/no need for monitoring.
    - No specific reversal agent for most; however, Andexanet alfa is available for factor Xa inhibitors.

# 2. Oral Anticoagulants

# A. Indirect Anticoagulants

- 1. Vitamin K Antagonists (VKAs)
  - **Example**: Warfarin (Coumadin)
  - Indications:
    - Atrial fibrillation (AF) to prevent stroke
    - Treatment and prophylaxis of DVT and PE
    - Mechanical heart valve management
  - Contraindications:
    - Active bleeding

- Pregnancy (can cause teratogenic effects)
- Notes:
  - Requires regular INR monitoring.
  - Affected by dietary vitamin K; patients need dietary counseling.

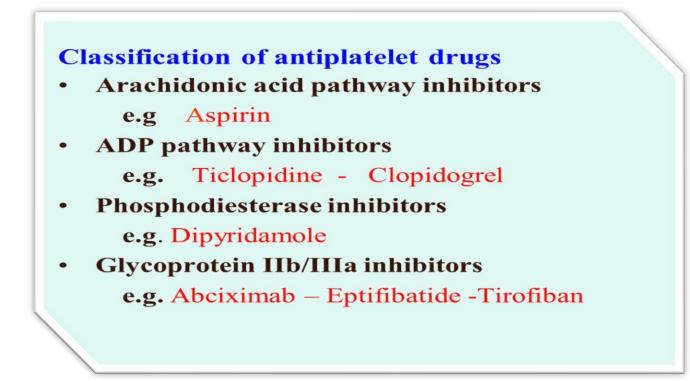
#### **B. Direct Anticoagulants**

- 1. Thrombin Inhibitors
  - Example: Dabigatran
  - Indications:
    - o Prevention of stroke and systemic embolism in AF
    - Treatment of DVT and PE
  - Contraindications:
    - Active bleeding
    - Mechanical heart valves (not recommended)
  - Notes:
    - An antidote, idarucizumab, is available for emergency reversal.

#### 2. Factor Xa Inhibitors

- Examples: Rivaroxaban, Apixaban, Edoxaban
- Indications:
  - Prevention of stroke in AF
  - Treatment and prevention of DVT and PE
- Contraindications:
  - Active bleeding
  - Severe renal impairment (especially caution for edoxaban)
- Notes:
  - Minimal monitoring required; however, should educate the patient about signs of bleeding.
  - Rivaroxaban is taken with food for effective absorption, while apixaban's timing is flexible







# **Classification of Antiplatelet Drugs**

- 1. Arachidonic Acid Pathway Inhibitors
  - Example: Aspirin
    - **Mechanism**: Inhibits cyclooxygenase-1 (COX-1), reducing thromboxane A2 production, which is essential for platelet aggregation.
    - Indications:
      - Primary and secondary prevention of cardiovascular events (e.g., myocardial infarction, stroke).
      - Acute coronary syndrome (ACS) management.
    - Contraindications:
      - Active gastrointestinal bleeding.
      - Severe hypersensitivity to aspirin or NSAIDs.
    - Clinical Notes:
      - Low-dose aspirin (75-100 mg daily) is effective for cardiovascular protection.

• Patients may require gastroprotective agents to mitigate gastrointestinal side effects.

#### 2. ADP Pathway Inhibitors

#### • Examples: Clopidogrel, Ticlopidine

- **Mechanism**: Inhibit the P2Y12 receptor on platelets, preventing activation and aggregation.
- Indications:
  - Clopidogrel: Used for secondary prevention of atherothrombotic events, ACS, and after PCI.
  - Ticlopidine: Less commonly used due to side effects; indicated for stroke prevention.

#### • Contraindications:

- Active bleeding.
- History of hypersensitivity to the drug.
- Clinical Notes:
  - Clopidogrel is a prodrug and may have variable efficacy due to genetic polymorphisms affecting metabolism.
  - Dual antiplatelet therapy (DAPT) with aspirin is often recommended after stent placement.

#### 3. Phosphodiesterase Inhibitors

#### • Example: Dipyridamole

- **Mechanism**: Inhibits phosphodiesterase, leading to increased levels of cyclic AMP in platelets, which inhibits aggregation.
- Indications:
  - Used in combination with aspirin for secondary prevention of stroke.
  - May be used in patients with peripheral vascular disease.
- **Contraindications**:
  - Active bleeding.
  - Severe hypotension or unstable angina.
- Clinical Notes:
  - Often used in combination with aspirin for enhanced antiplatelet effect.

#### 4. Glycoprotein IIb/IIIa Inhibitors

- Examples: Abciximab, Eptifibatide, Tirofiban
  - **Mechanism**: Block the glycoprotein IIb/IIIa receptor on activated platelets, preventing fibrinogen binding and platelet aggregation.
  - Indications:

• Used in high-risk patients undergoing PCI and in ACS management.

#### • Contraindications:

- Active bleeding.
- History of stroke or major surgery within the past 30 days.

#### • **Clinical Notes**:

- Abciximab has a longer duration of action and is typically used in a hospital setting.
- Eptifibatide and tirofiban are shorter-acting and can be used in outpatient settings.



# **Classification of Thrombolytic Drugs**

Generation of Thrombolytic Drug	Fibrin Specific	Non-Fibrin Specific
First Generation		Urokinase *
		Streptokinase *
Second Generation	Recombinant tissue plasminogen activator (t-PA) *	Pro-urokinase (scum-PA) *
	Alteplase	Sk-plasminogen activating complex (APSAC) *

# **Details of Thrombolytic Agents**

### **1. First Generation Thrombolytics**

Urokinase:

- **Mechanism**: Activates plasminogen to plasmin, leading to fibrin degradation.
- **Indications**: Used for the treatment of pulmonary embolism and acute myocardial infarction.
- **Notes**: Less fibrin-specific, which may lead to systemic fibrinolysis and increased bleeding risk.
- Streptokinase:
  - **Mechanism**: Forms a complex with plasminogen, converting it to plasmin.
  - **Indications**: Used in acute myocardial infarction and severe deep vein thrombosis.
  - **Notes**: Can induce allergic reactions; not fibrin-specific, leading to potential bleeding complications.

# 2. Second Generation Thrombolytics

- Recombinant Tissue Plasminogen Activator (t-PA):
  - Example: Alteplase
  - **Mechanism**: Fibrin-specific; preferentially activates plasminogen bound to fibrin in clots.
  - **Indications**: First-line treatment for acute ischemic stroke, myocardial infarction, and massive pulmonary embolism.
  - **Notes**: Lower risk of systemic bleeding compared to first-generation agents.
- Pro-Urokinase (scum-PA):
  - **Mechanism**: Similar to urokinase but with improved fibrin specificity.
  - **Indications**: Used in specific thrombotic conditions.
- Sk-Plasminogen Activating Complex (APSAC):
  - **Mechanism**: A complex that activates plasminogen to plasmin.
  - Indications: Used in acute myocardial infarction.

